This listing of claims will replace all prior versions, and listings, of claims in the

application:

1. (currently amended) An agent method for improving the blood stability of an

mammalian endogenous ligand in a mammal, which comprises administering to the

mammal an effective amount of an antibody that has an affinity to the endogenous ligand

but does not neutralize the same substantially.

2. (currently amended) The agent method of claim 1, wherein the improved blood

stability of the endogenous ligand results in the enhancement of receptor activity-

regulatory action thereof.

3. (currently amended) The agent method of claim 1, wherein the neutralizing activity of

the antibody is about 80% or less.

4. (currently amended) The agent method of claim 1, wherein the blood concentration of

the endogenous ligand becomes about twice or more compared to the case where the

antibody is not administered.

5. (currently amended) The agent method of claim 1, wherein the blood half-life of the

complex of the endogenous ligand and the antibody is about twice or more as that of the

endogenous ligand alone.

6. (currently amended) The agent method of claim 1, wherein the blood half-life of the

free endogenous ligand is about one week or less.

7. (currently amended) The agent method of claim 1, wherein the endogenous ligand is a

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peptidic compound.

8. (currently amended) The agent method of claim 7, wherein the endogenous ligand is

one against a G protein-coupled receptor.

9. (currently amended) The agent method of claim 8, wherein the endogenous ligand is

one belonging to secretin/glucagon super family.

10. (currently amended) The agent method of claim 9, wherein the endogenous ligand is

selected from the group consisting of GLP-1, calcitonin, PACAP, VIP and analogs

thereof.

11. (currently amended) The agent method of claim 8, wherein the endogenous ligand is

selected from the group consisting of LHRH, metastin, GPR7/GPR8 ligand, MSH,

ghrelin, apelin and analogs thereof.

12. (currently amended) The agent method of claim 7, wherein the endogenous ligand is

selected from the group consisting of EPO, TPO, insulin, interferon, growth hormone,

GM-CSF, leptin, adiponectin and analogs thereof.

13. (currently amended) The agent method of claim 7, wherein the endogenous ligand is

selected from the group consisting of ANP, BNP, CNP, betacellulin, betacellulin-δ4,

adrenomedullin and analogs thereof.

14. (currently amended) The agent method of claim 1, which is for the prophylaxis and/or

treatment of a disease in which an increased blood concentration and/or a prolonged

blood half-life of the endogenous ligand are/is effective for the prophylaxis and/or

treatment thereof.

15. (currently amended) The agent method of claim 14, wherein the disease is selected

from the group consisting of metabolic disease, bone and joint disease, cardiovascular

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disease, cranial nerve disease, infectious disease, cancer, blood disorder, urologic disease,

infertility/erectile dysfunction, deficient growth and immunodeficiency.

16. (currently amended) A method for the prophylaxis and/or treatment of a disease in a

mammal, wherein an increased blood concentration and/or a prolonged blood half-life of

an endogenous ligand are/is effective for the prophylaxis and/or treatment of the disease,

which method comprises administering to the mammal an effective amount of an

antibody that has an affinity to the endogenous ligand but does not neutralize the same

substantially, without administering a compound the same as or substantially the same as

the endogenous ligand, so as to increase the blood stability of the endogenous ligand,

thereby enhancing a receptor activity-regulatory action of the ligand.

17. (canceled) A use of an antibody that has an affinity for an endogenous ligand but does

not neutralize the same substantially for the manufacture of an agent for the prophylaxis

and/or treatment of a disease in which an increased blood concentration and/or a

prolonged blood half-life of the endogenous ligand are/is effective for the prophylaxis

and/or treatment thereof.

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